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(54) Title: METHODS AND COMPOSITIONS FOR DETERMINING ENZYMATIC ACTIVITY

(57) Abstract: The present invention comprises crystalline polyketide synthases, isolated non-native polyketide synthases having the structural coordinates of said crystalline polyketide synthases, and nucleic acids encoding such non-native polyketide synthases. Also disclosed are methods of predicting the activity and/or substrate specificity of putative polyketide synthase, methods of identifying potential polyketide synthase substrates, and methods of identifying potential polyketide synthase inhibitors.

INTERNATIONAL SEARCH REPORT

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A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C12N15/29 C12N15/54 C12N9/10 C12Q1/48

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C12N C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, EMBL, BIOSIS, MEDLINE, EMBASE

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	JEZ J M ET AL.: "Structure of chalcone synthase and the molecular basis of plant polyketide biosynthesis." FASEB JOURNAL. MEETING INFO.: ANNUAL MEETING OF THE AMERICAN SOCIETIES FOR EXPERIMENTAL BIOLOGY ON BIOCHEMISTRY AND MOLECULAR BIOLOGY 99 SAN FRANCISCO, CALIFORNIA, USA MAY 16-20, 1999, vol. 13, no. 7, 23 April 1999 (1999-04-23), page A1392 XP002158823	1,6,7, 12-14, 16,17, 22-39
Y	abstract 355	13,15, 18,22, 25, 29-31, 34-36

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☒ Further documents are listed in the continuation of box C.

☐ Patent family members are listed in annex.

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>DATABASE PDB 'Online! ID 1BI5, 22 June 1999 (1999-06-22) FERRER J-L ET AL.: "Chalcone synthase from Alfalfa" XP002158822 cited in the application</p>	<p>1,6,7, 12-14, 16,17, 22-39</p>
Y	<p>the whole document</p>	<p>13,15, 18,22, 25, 29-31, 34-36</p>
X	<p>--- JUNGHANS H ET AL.: "Stress responses in alfalfa (<i>Medicago sativa</i> L.). 15. Characterization and expression patterns of members of a subset of the chalcone synthase multigene family" PLANT. MOL. BIOL., vol. 22, no. 2, May 1993 (1993-05), pages 239-253, XP002158812 cited in the application Note: 99.7% aa sequence identity of CHS2 with SEQ ID NO:1 in in 389 aa overlap. abstract figures 1-3</p>	<p>1,7</p>
Y	<p>--- RAIBER S ET AL.: "Molecular and enzymatic characterization of two stilbene synthases from Eastern white pine (<i>Pinus strobus</i>). A single Arg/His difference determines the activity and the pH dependence of the enzymes." FEBS LETT., vol. 361, no. 2-3, 20 March 1995 (1995-03-20), pages 299-302, XP002158813 abstract figure 2</p>	<p>13,22, 29,31, 34,36</p>
Y	<p>--- HELARIUTTA Y ET AL.: "Chalcone synthase-like genes active during corolla development are differentially expressed and encode enzymes with different catalytic properties in <i>Gerbera hybrida</i> (<i>Asteraceae</i>)" PLANT MOLECULAR BIOLOGY, vol. 28, no. 1, April 1995 (1995-04), pages 47-60, XP002158814 abstract figure 1</p> <p>--- -/--</p>	<p>13,15, 22,29, 31,34,36</p>

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Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	ECKERMANN S ET AL.: "New pathway to polyketides in plants" NATURE, vol. 396, 26 November 1998 (1998-11-26), pages 387-390, XP002158815 cited in the application abstract page 389, left-hand column, line 50-53 ---	13,15, 22,29, 31,34,36
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Y	PREISIG-MÜLLER R ET AL.: "Plant polyketide synthases leading to stilbenoids have a domain catalyzing malonyl-CoA:CO ₂ exchange, malonyl-CoA decarboxylation, and covalent enzyme modification and a site for chain lengthening" BIOCHEMISTRY, vol. 36, no. 27, 8 July 1997 (1997-07-08), pages 8349-8358, XP002158817 abstract tables 1,4 ---	18,25, 30,31, 35,36
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In ational Application No

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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